

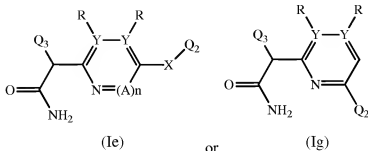
AMENDMENTS TO THE CLAIMS

Claims 3, 8-12, 15, 18-23, 25-29, and 34 are currently pending. Please amend claims 3 and 15, as indicated below. This listing of claims will replace all prior versions, and listings, of claims in the application.

Listing Of Claims

1-2. (Canceled)

3. (Currently amended) A compound of the formula:



wherein:

Q₃ is a 5-6 membered aromatic carbocyclic or heterocyclic ring system; or an 8-10 membered bicyclic ring system ~~comprising~~ consisting of aromatic carbocyclic rings, aromatic heterocyclic rings or a combination of an aromatic carbocyclic ring and an aromatic heterocyclic ring; wherein Q₃ is substituted with 1 to 4 substituents, each of which is independently selected from halo; C₁-C₃ alkyl optionally substituted with NR'₂, OR', CO₂R' or CONR'₂; O-(C₁-C₃)-alkyl optionally substituted with NR'₂, OR', CO₂R' or CONR'₂; NR'₂; OCF₃; CF₃; NO₂; CO₂R'; CONHR'; SR'; S(O₂)N(R')₂; SCF₃; CN; N(R')C(O)R⁴; N(R')C(O)OR⁴; N(R')C(O)C(O)R⁴; N(R')S(O₂)R⁴; N(R')R⁴; N(R⁴)₂; OR⁴; OC(O)R⁴; OP(O)₃H₂; or N=CH-N(R')₂;

Q₂ is selected from 5-6 membered aromatic carbocyclic or heterocyclic ring systems, or 8-10 membered bicyclic ring systems consisting of aromatic carbocyclic rings, aromatic heterocyclic rings or a combination of an aromatic carbocyclic ring and an aromatic heterocyclic ring; wherein:

Q₂ is optionally substituted with up to 4 substituents, independently selected from halo, CH=N-OH, or CH=O; C₁-C₃ straight or branched alkyl optionally substituted with NR'₂, OR', CO₂R', S(O₂)N(R')₂, N=CH-N(R')₂, R³, NH-CH₃, NHCH₂CH₂OH, NHCH₂CH(OH)CH₂OH, CH₂OCH₂OCH₃, NHCH₂CH₂NH₂, NH-phenyl, piperazinyl, pyrrolidinyl or CONR'₂; O-(C₁-C₃)-alkyl optionally substituted with NR'₂, OR', CO₂R', S(O₂)N(R')₂, N=CH-N(R')₂, R³, or CONR'₂; NR'₂; OCF₃; CF₃; NO₂; CO₂R'; CONHR'; R³; OR³; NHR³; SR³; C(O)R³; C(O)N(R')R³; C(O)OR³; SR'; S(O₂)N(R')₂; SCF₃; N=CH-N(R')₂; CH=N-OH; CH=O; or CN;

wherein R' is selected from hydrogen, (C₁-C₃)-alkyl; (C₂-C₃)-alkenyl or alkynyl; phenyl or phenyl substituted with 1 to 3 substituents independently selected from halo, methoxy, cyano, nitro, amino, hydroxy, methyl or ethyl;

R³ is selected from a 5-6 membered aromatic carbocyclic or heterocyclic ring system;

R⁴ is (C₁-C₄)-alkyl optionally substituted with N(R')₂, OR', CO₂R', CON(R')₂, or SO₂N(R')₂; or a 5-6 membered carbocyclic or heterocyclic ring system optionally substituted with N(R')₂, OR', CO₂R', CON(R')₂, or SO₂N(R')₂;

X is selected from -S-, -O-, -S(O₂)-, -S(O)-, -N(R²)-, -N(R²)-S(O₂)-, -N(R²)-C(O)O-, -O-C(O)-N(R²), -C(O)-, -C(O)O-, -O-C(O)-, -C(O)-N(R²)-, -N(R²)-C(O)-, -C(R²)₂-, -C(OR²)₂-, -CH(OH)-;

each R is independently selected from hydrogen, -R², -N(R²)₂, -OR², SR², -C(O)-N(R²)₂, -S(O₂)-N(R²)₂, or -C(O)-OR², wherein two adjacent R are optionally bound to one another and, together with each carbon to which they are respectively bound, form a 4-8 membered carbocyclic or heterocyclic ring;

R² is selected from hydrogen, (C₁-C₃)-alkyl, or (C₂-C₃)-alkenyl; each optionally substituted with -N(R')₂, -OR', SR', -C(O)-N(R')₂, -S(O₂)-N(R')₂, -C(O)-OR', or R³;

Y is C;

A, if present, is CR'; and

n is 1;

provided that when a compound is of formula Ig, Q₃ is 2,6-dichlorophenyl and both R substituents are H, then Q₂ is neither phenyl nor p-fluorophenyl; and

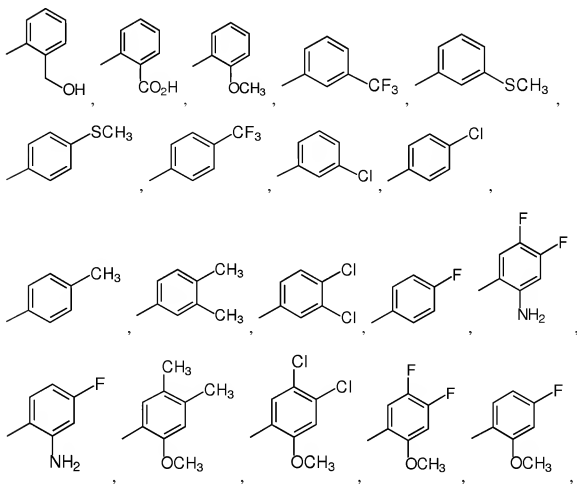
when a compound is of formula Ie, and Q₃ is 2,6-dichlorophenyl, both R substituents are H, and X is S, then Q₂ is not phenyl.

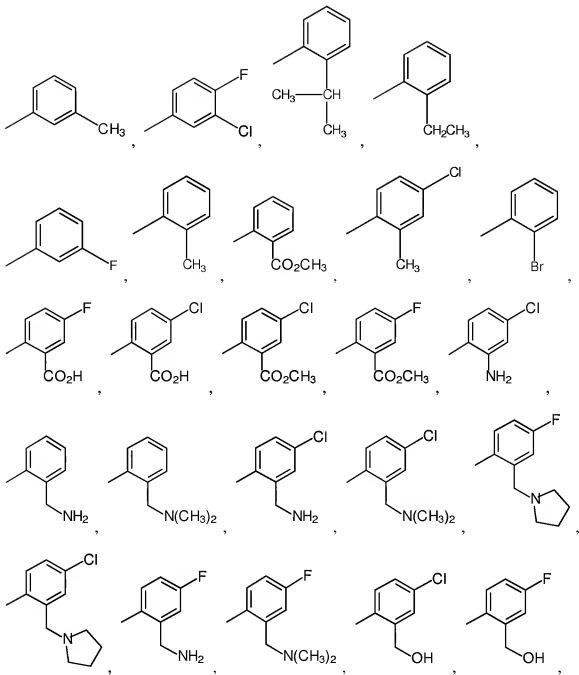
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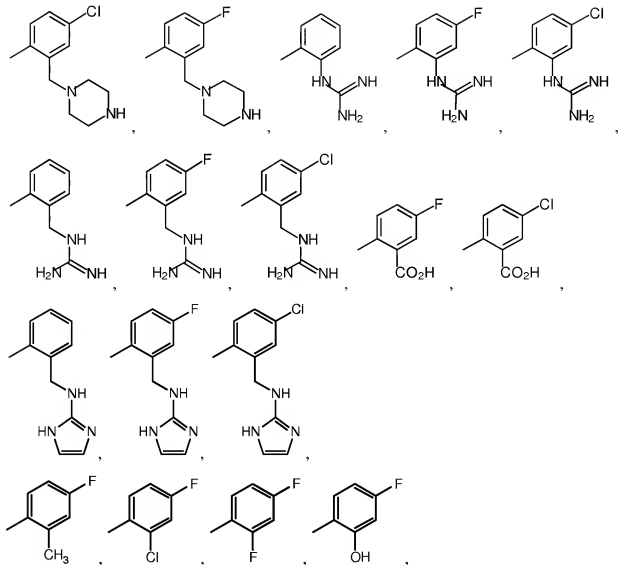
8. (Previously presented) The compound according to claim 3, wherein Q₂ is selected from phenyl or pyridyl and wherein Q₂ optionally contains up to 3 substituents, each of which is independently selected from chloro, fluoro, bromo, methyl, ethyl, isopropyl, -

OCH₃, -OH, -NH₂, -CF₃, -OCF₃, -SCH₃, -OCH₃, -C(O)OH, -C(O)OCH₃, -CH₂NH₂, -N(CH₃)₂,
 -CH₂-pyrrolidine and -CH₂OH.

9. (Previously presented) The compound according to claim 8, wherein, Q₂ is selected from:







unsubstituted 2-pyridyl or unsubstituted phenyl.

10. (Previously presented) The compound according to claim 9, wherein Q_2 is selected from phenyl, 2-isopropylphenyl, 3,4-dimethylphenyl, 2-ethylphenyl, 3-fluorophenyl, 2-methylphenyl, 3-chloro-4-fluorophenyl, 3-chlorophenyl, 2-carbomethoxyphenyl, 2-carboxyphenyl, 2-methyl-4-chlorophenyl, 2-bromophenyl,

2-pyridyl, 2-methylenedihydroxyphenyl, 4-fluorophenyl, 2-methyl-4-fluorophenyl,
2-chloro-4-fluorophenyl, 2,4-difluorophenyl, 2-hydroxy-4-fluorophenyl or
2-methylenedihydroxy-4-fluorophenyl.

11. (Previously presented) The compound according to claim 3, wherein X
is selected from $-S-$, $-O-$, $-S(O_2)-$, $-S(O)-$, $-N(R^2)-$,
 $-C(R^2)_2-$ or $-C(O)-$.

12. (Previously presented) The compound according to claim 11, wherein X
is S.

13-14. (Canceled)

15. (Currently amended) The compound according to claim[[14]]3,
wherein each R attached to Y is independently selected from hydrogen or methyl.

16-17. (Canceled)

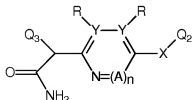
18. (Previously presented) The compound according to claim 3, wherein Q_3
is substituted with 2 to 4 substituents, wherein at least one of said substituents is present in the
ortho position relative to the point of attachment of Q_3 to the rest of the inhibitor.

19. (Original) The compound according to claim 18, wherein both ortho
positions are occupied by one of said independently selected substituents.

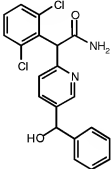
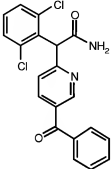
20. (Original) The compound according to claim 19, wherein Q_3 is a monocyclic carbocyclic ring; and each of said ortho substituents on Q_3 are independently selected from halo or methyl.

21. (Previously presented) The compound according to claim 19, wherein Q_3 contains 1 to 2 substituents in addition to said ortho substituents, said additional substituents being independently selected from NR'^2 , OR' , CO_2R' , CN , $N(R')C(O)R^4$, $N(R')C(O)OR^4$, $N(R')C(O)C(O)R^4$, $N(R')S(O_2)R^4$, $N(R')R^4$, $N(R^4)_2$, OR^4 , $OC(O)R^4$, $OP(O)_3H_2$; or $N=CH-N(R')_2$.

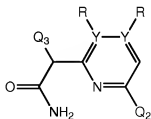
22. (Previously presented) The compound according to claim 3, wherein said compound is a compound of formula Ie:



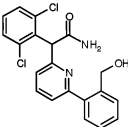
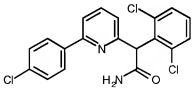
and is selected from any one of the following compounds:

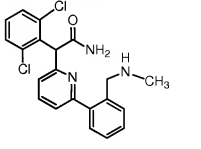
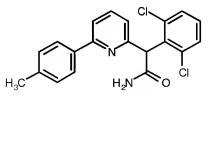
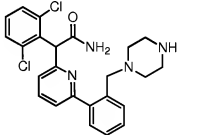
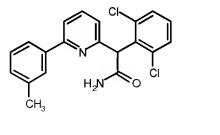
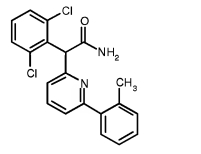
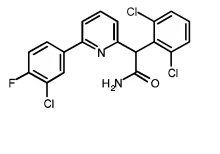
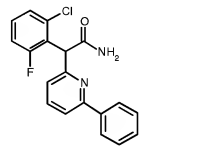
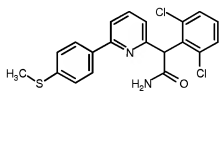
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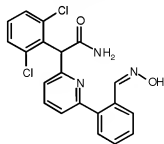
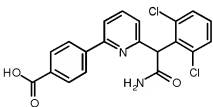
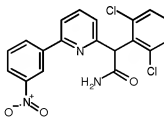
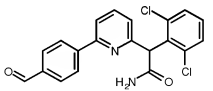
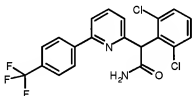
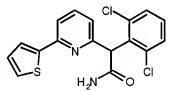
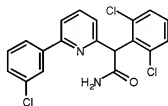
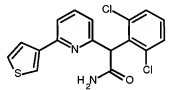
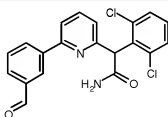
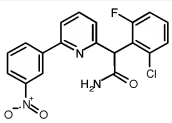
23. (Previously presented) The compound according to claim 3, wherein said compound is a compound of formula Ig:

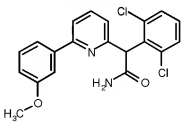
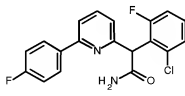
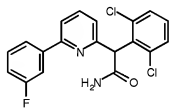
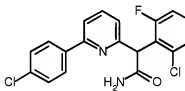
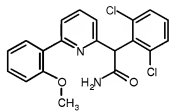
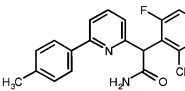
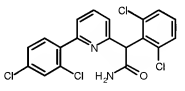
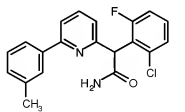
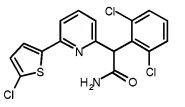
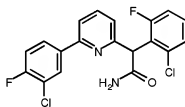


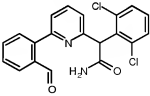
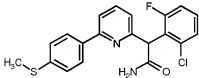
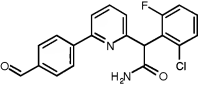
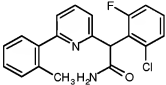
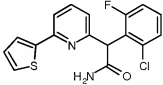
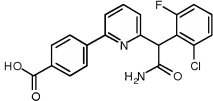
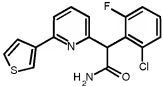
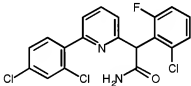
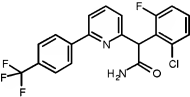
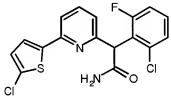
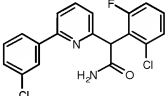
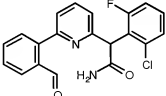
and is selected from any one of the following compounds:

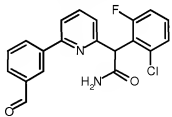
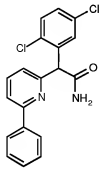
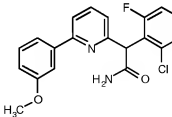
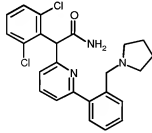
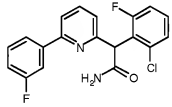
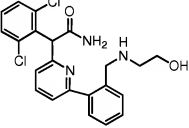
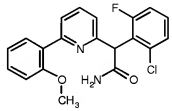
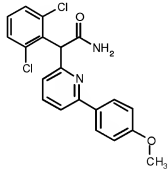
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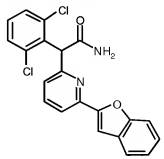
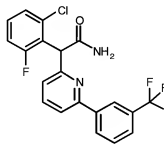
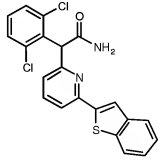
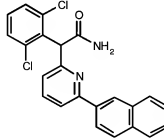
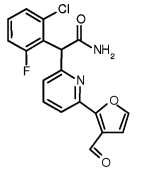
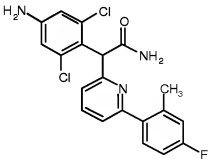
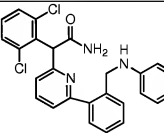
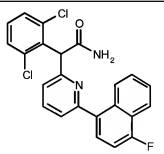
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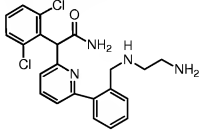
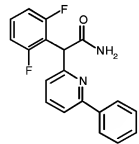
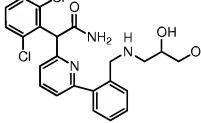
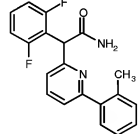
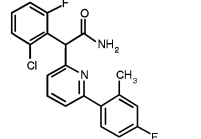
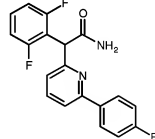
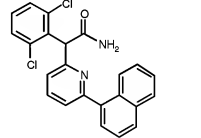
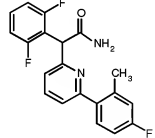
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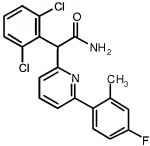
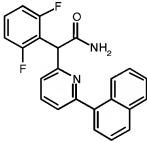
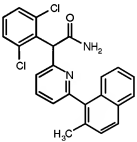
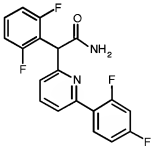
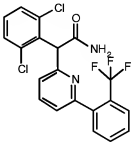
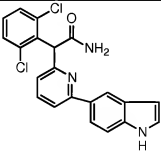
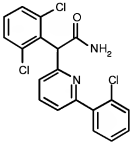
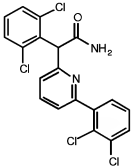
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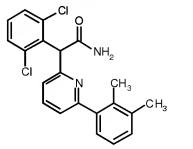
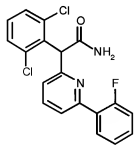
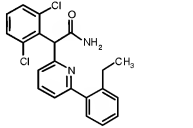
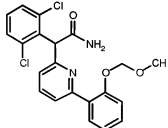
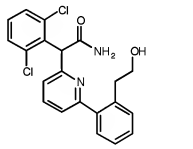
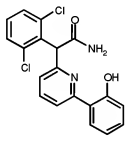
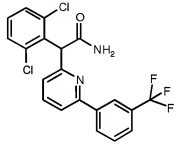
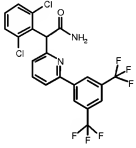
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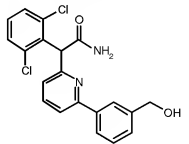
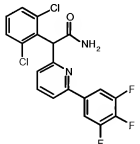
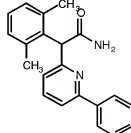
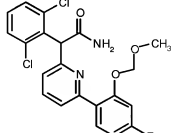
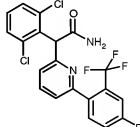
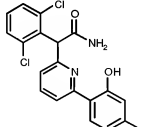
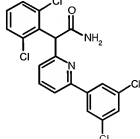
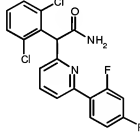
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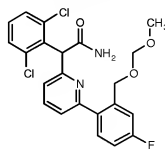
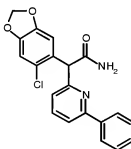
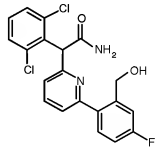
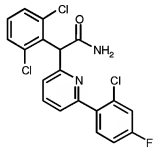
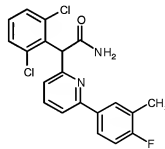
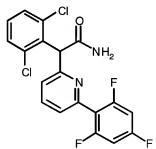
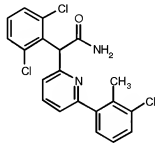
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24. (Canceled)

25. (Previously presented) A pharmaceutical composition comprising an amount of a compound according to claim 3 effective to inhibit p38, and a pharmaceutically acceptable carrier.

26. (Previously presented) A method of treating inflammatory diseases, destructive bone disorders, reperfusion/ischemia in stroke, myocardial ischemia, renal ischemia, cardiac hypertrophy, rheumatoid arthritis, inflammatory bowel disease, ulcerative colitis, or Crohn's disease in a patient, said method comprising administering to said patient a composition according to claim 25.

27. (Previously presented) The method according to claim 26, wherein said method is used to treat an inflammatory disease selected from acute pancreatitis, chronic pancreatitis, asthma, allergies, or adult respiratory distress syndrome.

28. (Previously presented) The method according to claim 26, wherein said method is used to treat rheumatoid arthritis, inflammatory bowel disease, ulcerative colitis, or Crohn's disease.

29. (Previously presented) The method according to claim 26, wherein said method is used to treat a destructive bone disorder selected from osteoarthritis, osteoporosis or multiple myeloma-related bone disorder.

30-33. (Canceled)

34. (Previously presented) The method according to claim 26, wherein said method is used to treat ischemia/reperfusion in stroke, myocardial ischemia, or renal ischemia.

35-37. (Canceled)